

=> d his

(FILE 'HOME' ENTERED AT 06:23:56 ON 28 OCT 2004)  
SET COST OFF

FILE 'HCAPLUS' ENTERED AT 06:27:06 ON 28 OCT 2004

L1 2 S US20040127702/PN OR (US2003-666175# OR US2002-425596# OR US20  
E PEARLMAN B/AU  
L2 31 S E4,E7-E9  
E ANTOSZ F/AU  
L3 16 S E3-E5  
L4 183 S ?EPLERENON?

FILE 'REGISTRY' ENTERED AT 06:28:49 ON 28 OCT 2004

L5 1 S 107724-20-9

FILE 'HCAPLUS' ENTERED AT 06:29:31 ON 28 OCT 2004

SET SMARTSELECT ON  
L6 SEL L1 1- RN : 100 TERMS  
SET SMARTSELECT OFF

FILE 'REGISTRY' ENTERED AT 06:29:31 ON 28 OCT 2004

L7 100 S L6  
L8 27 S L7 AND OC4-C5-C6-C6-C6/ES  
L9 6 S L8 AND OC4/ES  
L10 1 S L9 AND C27H32O4  
E C27H32O4/MF  
L11 1 S E3 AND OC4-C5-C6-C6-C6/ES AND OC4/ES  
L12 1 S L10,L11  
SEL RN  
L13 0 S E1/CRN

FILE 'HCAOLD' ENTERED AT 06:32:59 ON 28 OCT 2004

L14 0 S L12

FILE 'USPATFULL, USPAT2' ENTERED AT 06:33:00 ON 28 OCT 2004

L15 1 S L12

FILE 'HCAPLUS' ENTERED AT 06:33:01 ON 28 OCT 2004

L16 3 S L12  
L17 1 S L16 AND L1-L3  
L18 2 S L16 AND (PHARMACIA? OR UPJOHN?)/PA,CS  
L19 3 S L16-L18

FILE 'REGISTRY' ENTERED AT 06:37:28 ON 28 OCT 2004

L20 STR  
L21 0 S L20  
L22 STR L20  
L23 7 S L22  
L24 2101 S L22 FUL  
SAV TEMP L24 QAZI666/A  
L25 45 S L20 FUL SUB=L24  
SAV TEMP L25 QAZI666A/A  
L26 STR L20  
L27 12 S L26 FUL SUB=L25  
SAV L27 TEMP QAZI666B/A  
L28 11 S L27 NOT L12

FILE 'HCAPLUS' ENTERED AT 06:40:24 ON 28 OCT 2004

L29 2 S L28  
L30 1 S L29 AND L1-L3  
L31 1 S L29 AND (PHARMACIA? OR UPJOHN?)/PA,CS  
L32 3 S L19,L29-L31

FILE 'USPATFULL, USPAT2' ENTERED AT 06:40:50 ON 28 OCT 2004

L33 1 S L28  
L34 1 S L15,L33

=> fil reg

FILE 'REGISTRY' ENTERED AT 06:41:14 ON 28 OCT 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 26 OCT 2004 HIGHEST RN 769912-90-5

DICTIONARY FILE UPDATES: 26 OCT 2004 HIGHEST RN 769912-90-5

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

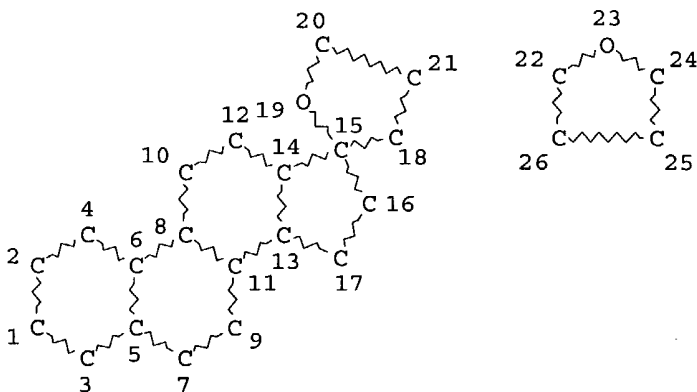
Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at:

<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> d sta que 127

L20 STR



NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

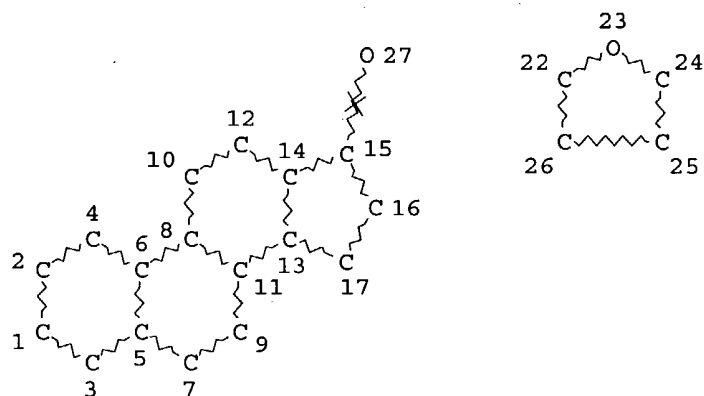
GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 26

STEREO ATTRIBUTES: NONE

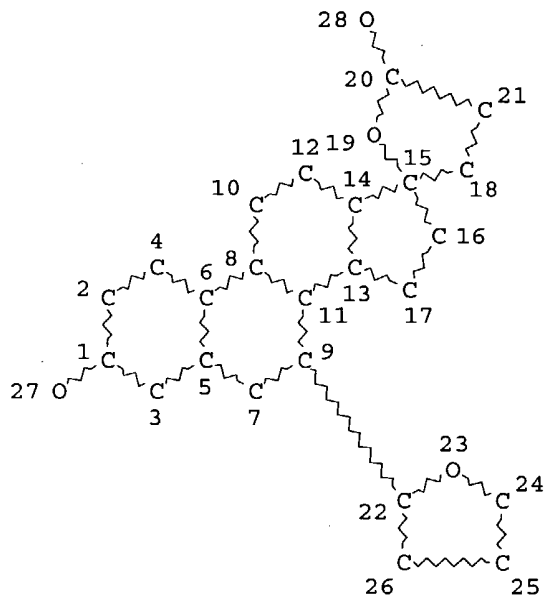
L22 STR



NODE ATTRIBUTES:  
 DEFAULT MLEVEL IS ATOM  
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
 RING(S) ARE ISOLATED OR EMBEDDED  
 NUMBER OF NODES IS 23

STEREO ATTRIBUTES: NONE  
 L24 2101 SEA FILE=REGISTRY SSS FUL L22  
 L25 45 SEA FILE=REGISTRY SUB=L24 SSS FUL L20  
 L26 STR



NODE ATTRIBUTES:  
 DEFAULT MLEVEL IS ATOM  
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
 RING(S) ARE ISOLATED OR EMBEDDED  
 NUMBER OF NODES IS 28

STEREO ATTRIBUTES: NONE  
 L27 12 SEA FILE=REGISTRY SUB=L25 SSS FUL L26

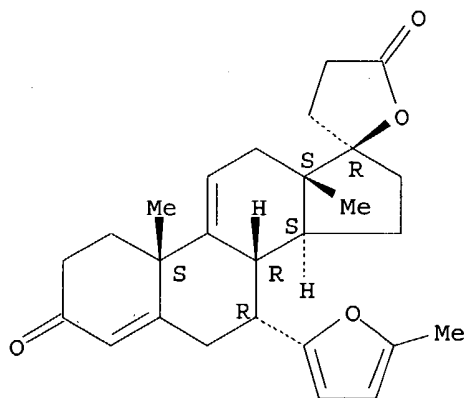
100.0% PROCESSED 13 ITERATIONS  
SEARCH TIME: 00.00.01

12 ANSWERS

=&gt; d ide can l12

L12 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-40-5 REGISTRY  
CN Pregna-4,9(11)-diene-21-carboxylic acid, 17-hydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H32 O4  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT, USPATFULL  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

Absolute stereochemistry.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

2 REFERENCES IN FILE CA (1907 TO DATE)  
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

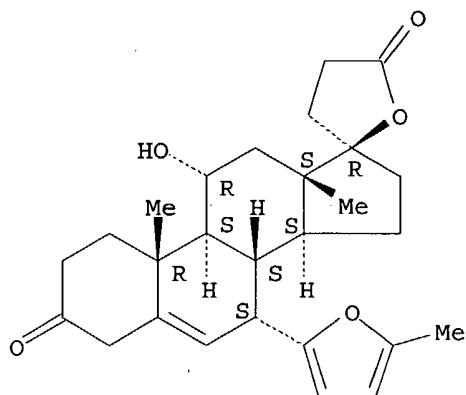
REFERENCE 1: 140:423864

REFERENCE 2: 139:307925

=&gt; d ide can tot l28

L28 ANSWER 1 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 690958-98-6 REGISTRY  
CN Pregn-5-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H34 O5  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT, USPATFULL  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)

Absolute stereochemistry.



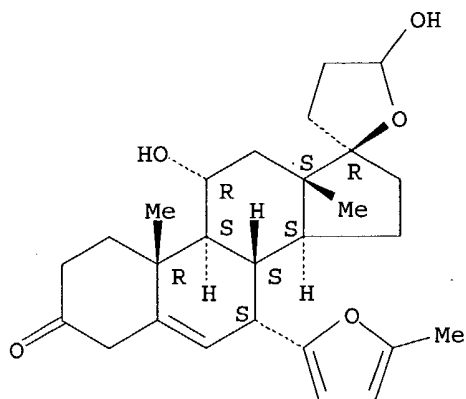
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:423864

L28 ANSWER 2 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 690958-89-5 REGISTRY  
CN Spiro[androst-5-ene-17,2'-(3'H)-furan]-3-one, 4',5'-dihydro-5',11-dihydroxy-7-(5-methyl-2-furanyl)-, (7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )-(9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H36 O5  
SR CA  
LC STN Files: CA, CAPLUS, USPATFULL  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)

Absolute stereochemistry.



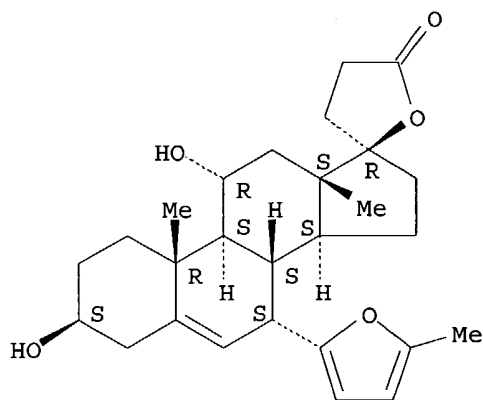
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:423864

L28 ANSWER 3 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
 RN 690958-83-9 REGISTRY  
 CN Pregn-5-ene-21-carboxylic acid, 3,11,17-trihydroxy-7-(5-methyl-2-furanyl)-  
 ,  $\gamma$ -lactone, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA  
 INDEX NAME)  
 FS STEREOSEARCH  
 MF C27 H36 O5  
 SR CA  
 LC STN Files: CA, CAPLUS, CASREACT, USPATFULL  
 DT.CA Caplus document type: Patent  
 RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT  
 (Reactant or reagent)

Absolute stereochemistry.



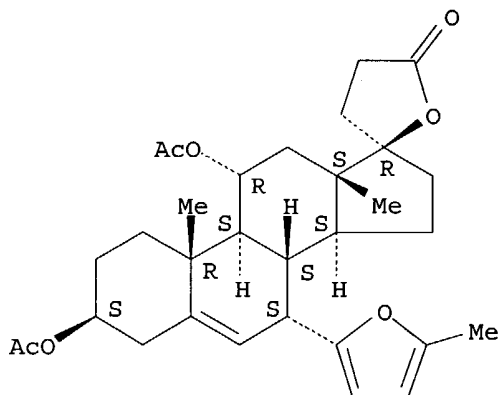
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:423864

L28 ANSWER 4 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
 RN 690958-80-6 REGISTRY  
 CN Spiro[androst-5-ene-17,2'(5'H)-furan]-5'-one, 3,11-bis(acetyloxy)-3',4'-  
 dihydro-7-(5-methyl-2-furanyl)-, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )-  
 (9CI) (CA INDEX NAME)  
 FS STEREOSEARCH  
 MF C31 H40 O7  
 SR CA  
 LC STN Files: CA, CAPLUS, CASREACT, USPATFULL  
 DT.CA Caplus document type: Patent  
 RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT  
 (Reactant or reagent)

Absolute stereochemistry.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:423864

L28 ANSWER 5 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN

RN 690958-45-3 REGISTRY

CN Spiro[androst-5-ene-17,2'(3'H)-furan]-3,5',11-triol, 4',5'-dihydro-7-(5-methyl-2-furanyl)-, (3β,7α,11α,17β)- (9CI) (CA INDEX NAME)

FS STEREOSEARCH

MF C27 H38 O5

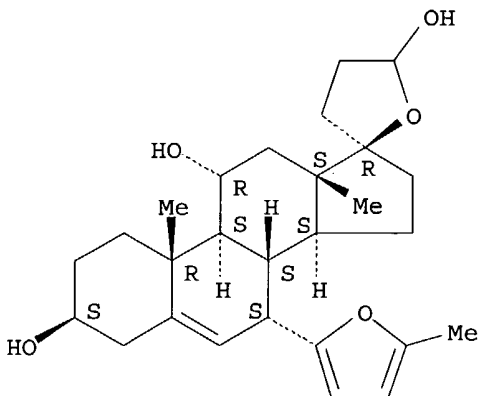
SR CA

LC STN Files: CA, CAPLUS, CASREACT, USPATFULL

DT.CA Caplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)

Absolute stereochemistry.



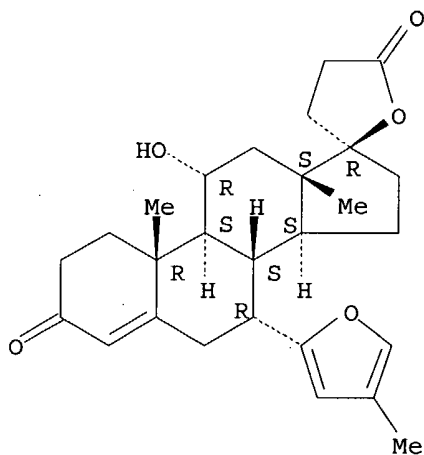
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:423864

L28 ANSWER 6 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-54-1 REGISTRY  
CN Pregn-4-ene-21-carboxylic acid, 11,17-dihydroxy-7-(4-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H34 O5  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

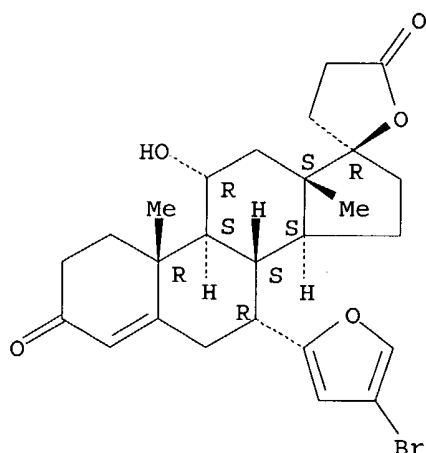
1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 139:307925

L28 ANSWER 7 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-53-0 REGISTRY  
CN Pregn-4-ene-21-carboxylic acid, 7-(4-bromo-2-furanyl)-11,17-dihydroxy-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C26 H31 Br O5  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.





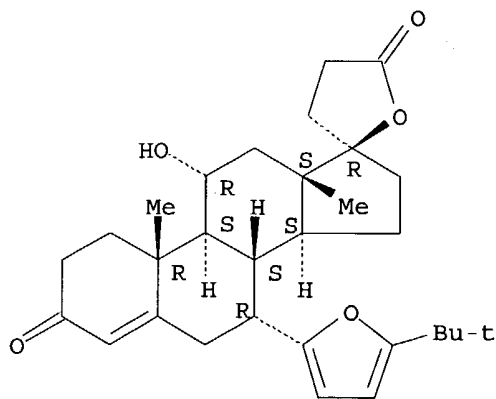
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 139:307925

L28 ANSWER 8 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-52-9 REGISTRY  
CN Pregn-4-ene-21-carboxylic acid, 7-[5-(1,1-dimethylethyl)-2-furanyl]-11,17-dihydroxy-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI)  
(CA INDEX NAME)  
FS STEREOSEARCH  
MF C30 H40 O5  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.



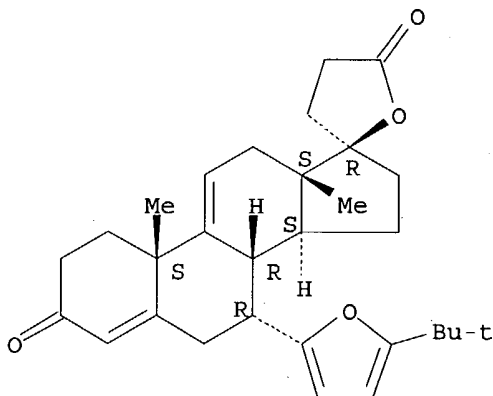
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 139:307925

L28 ANSWER 9 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-51-8 REGISTRY  
CN Pregna-4,9(11)-diene-21-carboxylic acid, 7-[5-(1,1-dimethylethyl)-2-furanyl]-17-hydroxy-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ )- (9CI)  
(CA INDEX NAME)  
FS STEREOSEARCH  
MF C30 H38 O4  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.



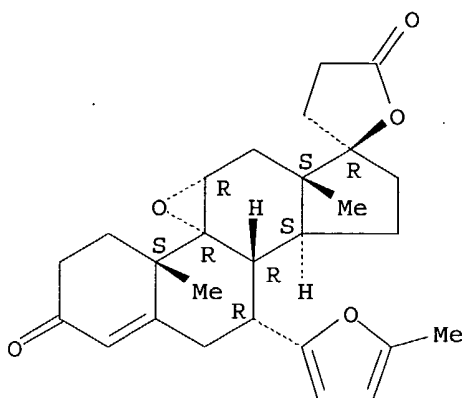
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 139:307925

L28 ANSWER 10 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-48-3 REGISTRY  
CN Pregn-4-ene-21-carboxylic acid, 9,11-epoxy-17-hydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI)  
(CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H32 O5  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT  
DT.CA Caplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.



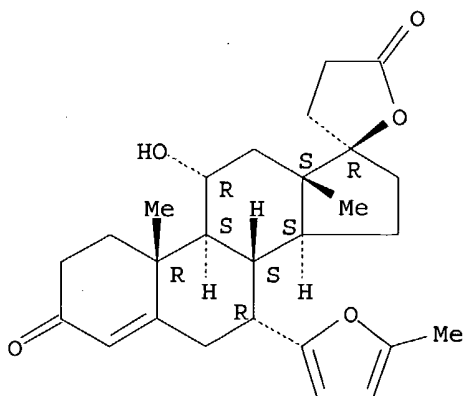
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1907 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 139:307925

L28 ANSWER 11 OF 11 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 610785-47-2 REGISTRY  
CN Pregn-4-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H34 O5  
SR CA  
LC STN Files: CA, CAPLUS, CASREACT, USPATFULL  
DT.CA CAplus document type: Patent  
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

Absolute stereochemistry.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

2 REFERENCES IN FILE CA (1907 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 140:423864

REFERENCE 2: 139:307925

=&gt; fil hcaplus

FILE 'HCAPLUS' ENTERED AT 06:41:48 ON 28 OCT 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 28 Oct 2004 VOL 141 ISS 18

FILE LAST UPDATED: 27 Oct 2004 (20041027/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=&gt; d l32 all hitstr tot

L32 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:817907 HCAPLUS

ED Entered STN: 07 Oct 2004

TI Preparation of spirosteroids from 17-alkenyl or 17-alkynyl substrate via carbonylation, hydrogenation, dehydrogenation, furylation and other transformations

IN Franczyk, Thaddeus S., II; Wagner, Grace M.

PA **Pharmacia Corporation, USA**

SO PCT Int. Appl., 177 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM C07J

CC 32-1 (Steroids)

FAN.CNT 1

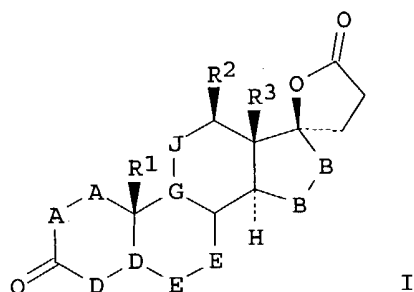
|    | PATENT NO.   | KIND | DATE     | APPLICATION NO. | DATE     |
|----|--|------|----------|-----------------|----------|
| PI | WO 2004085458  | A2   | 20041007 | WO 2004-US8629  | 20040322 |
|    | W:   |      |          |                 |          |
|    | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW |      |          |                 |          |
|    | RW:  |      |          |                 |          |
|    | BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |      |          |                 |          |

PRAI US 2003-456716P P 20030321

CLASS

| PATENT NO. | CLASS | PATENT FAMILY CLASSIFICATION CODES |
|------------|-------|------------------------------------|
|------------|-------|------------------------------------|

WO 2004085458 ICM C07J  
GI



- AB Steroids such as I (R1, R2, R3 = H, halo, haloalkyl, OH, alkyl, alkoxy, hydroxyalkyl, alkoxyalkyl, hydroxycarbonyl, CN, aryloxy; A-A, B-B, D-D, and G-J = substituted or unsubstituted double or single bond with R groups similar to those for R1, R2, R3 in the substituted case) comprising a 17-spirolactone or corresponding open lactone structure is obtained by carbonylation of a 17-alkenyl or 17-alkynyl substrate. A 17-alkenyl intermediate may be prepared by semi-hydrogenation of a 17-alkynyl group. Multiple reaction schemes are disclosed for preparation of a 3-keto-9,11-epoxy-17-spirolactone steroid such as eplerenone. Novel intermediates are also disclosed, as well as steps for forming such novel intermediates, or converting them to further intermediates or products, by semi-hydrogenation, carbonylation, 6,7-dehydrogenation, furylation or other transformations or combinations thereof.
- ST spiro-lactone steroid eplerenone prepn; hydrogenation carbonylation dehydrogenation furylation spiro-lactone steroid prepn
- IT Carbonylation  
Dehydrogenation  
Hydrogenation  
(preparation of spiro-lactone steroid derivs. from 17-alkenyl or 17 alkynyl steroids via carbonylation, semi-hydrogenation, dehydrogenation, furylation and other transformations)
- IT Steroids  
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)  
(spiro; preparation of spiro-lactone steroid derivs. from 17-alkenyl or 17 alkynyl steroids via carbonylation, semi-hydrogenation, dehydrogenation, furylation and other transformations)
- IT 192704-66-8P 610269-44-8P 610269-45-9P 610269-50-6P 610269-51-7P  
769146-02-3P 769146-04-5P  
RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of spiro-lactone steroid derivs. from 17-alkenyl or 17 alkynyl steroids via carbonylation, semi-hydrogenation, dehydrogenation, furylation and other transformations)
- IT 41850-21-9P 95716-71-5P 107724-20-9P, Eplerenone 610785-40-5P  
769146-08-9P  
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)  
(preparation of spiro-lactone steroid derivs. from 17-alkenyl or 17 alkynyl steroids via carbonylation, semi-hydrogenation, dehydrogenation, furylation and other transformations)
- IT 434-03-7, Ethisterone 534-22-5 28319-72-4 95716-70-4 769146-05-6  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(preparation of spiro-lactone steroid derivs. from 17-alkenyl or 17 alkynyl steroids via carbonylation, semi-hydrogenation, dehydrogenation, furylation and other transformations)

IT 610785-40-5P

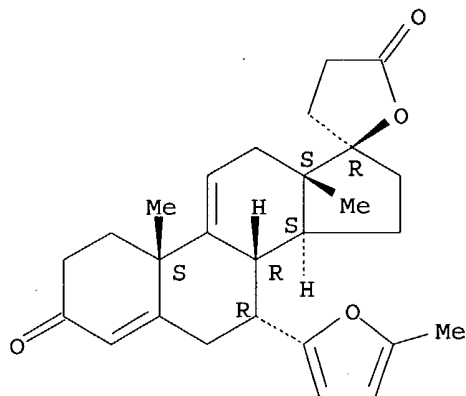
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(preparation of spirolactone steroid derivs. from 17-alkenyl or 17 alkynyl steroids via carbonylation, semi-hydrogenation, dehydrogenation, furylation and other transformations)

RN 610785-40-5 HCAPLUS

CN Pregna-4,9(11)-diene-21-carboxylic acid, 17-hydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L32 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:414628 HCAPLUS

DN 140:423864

ED Entered STN: 21 May 2004

TI Processes for preparing C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one

IN Wuts, Peter Guillaume Marie

PA USA

SO U.S. Pat. Appl. Publ., 23 pp.

CODEN: USXXCO

DT Patent

LA English

IC ICM A61K031-58

ICS C07J043-00; C07J021-00; C07J017-00

NCL 514173000; 514176000; 514178000; 540041000; 540107000; 540114000

CC 32-4 (Steroids)

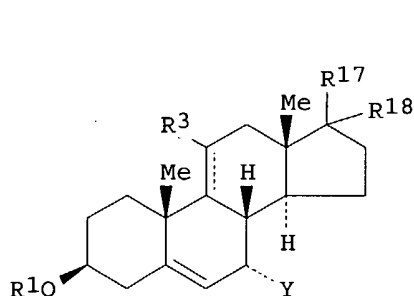
Section cross-reference(s): 16

FAN.CNT 1

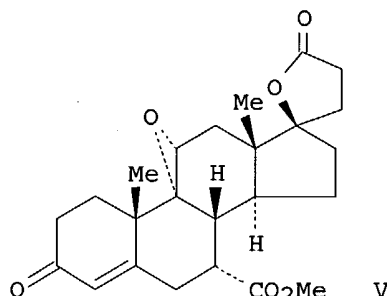
| PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE     |
|---|------|----------|-----------------|----------|
| US 2004097475   | A1   | 20040520 | US 2003-392945  | 20030321 |
| WO 2004043986   | A1   | 20040527 | WO 2003-US7284  | 20030321 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |      |          |                 |          |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG  |      |          |                 |          |
| PRAI US 2002-424488P  | P    | 20021107 |                 |          |

## CLASS

| PATENT NO.    | CLASS    | PATENT FAMILY CLASSIFICATION CODES                               |
|---------------|----------|--|
| US 2004097475 | ICM      | A61K031-58   |
|               | ICS      | C07J043-00; C07J021-00; C07J017-00                               |
|               | NCL      | 514173000; 514176000; 514178000; 540041000; 540107000; 540114000 |
| OS            | CASREACT | 140:423864; MARPAT 140:423864                                    |
| GI            |          |  |



II



V

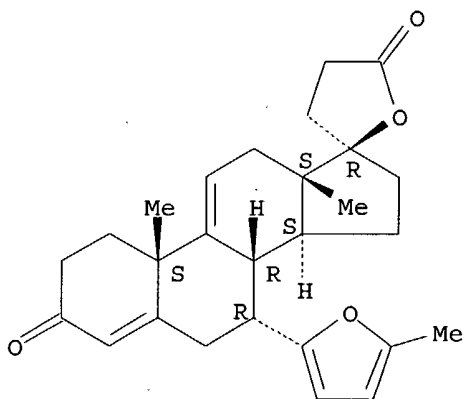
- AB The present invention discloses a process for the transformation of 5-androsten-3 $\beta$ -ol-17-one (I) to C-7 substituted steroids, such as II [R1 = H, COR2; R2 = alkyl, alkoxy; R3 = H, OR1; R17R18 = O, lactone; Y = CN, CH2CH:CH2, 5-(C1-6-alkyl)-2-furyl, 1-(C1-6-alkyl)-2-pyrrolyl, CHR4C(O)aryl, CHR4C(O)alkyl, CHR4C(O)X-aryl, CHR4C(O)X-alkyl; R4 = alkyl, aryl; X = O, S, dashed bond = single bond or double bond]. Thus, bioconversion of I to 5-androsten-3 $\beta$ ,7 $\beta$ -diol-17-one (III) was performed using a submerged culture of *Diplodia gossypina* ATCC 20571. III was subsequently converted to 5-androsten-3 $\beta$ ,7 $\beta$ ,11 $\alpha$ -triol-17-one (IV) using a submerged culture of *Aspergillus ochraceus* ATCC 18500. IV can also be obtained from II using a submerged culture of *Absidia coerulea* ATCC 6647. These intermediates are useful in the preparation of eplerenone (V).
- ST carboxysteroid eplerenone prepn androstenol deriv hydroxylation androstenone; hydroxyandrostenone dihydroxyandrostenone trihydroxyandrostenone prepn
- IT *Aspergillus ochraceus*  
(ATCC 18500; for 11 $\alpha$ -hydroxylation of 5-androsten-3 $\beta$ ,7 $\beta$ -diol-17-one in preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT *Diplodia gossypina*  
(ATCC 20571; for 7 $\beta$ -hydroxylation of 5-androsten-3 $\beta$ -ol-17-one in preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT *Absidia coerulea*  
(ATCC 6647; for 7 $\beta$ ,11 $\alpha$ -dihydroxylation of 5-androsten-3 $\beta$ -ol-17-one in preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Hydroxylation  
(biol.; during preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Lewis acids  
RL: CAT (Catalyst use); USES (Uses)  
(catalyst; during preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Bromination  
Dehydration reaction  
Ozonization

- (during preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Isomerization  
(of 4,5-double bond during preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Nucleophiles  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Steroids, preparation  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Sterols  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Acylation  
Esterification  
Hydroformylation  
Oxidation  
Silylation  
(stereoselective; during preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT Allylation  
(stereoselective; preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 537718-07-3P  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); RCT (Reactant); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 95716-70-4P 192704-56-6P 537718-14-2P **610785-40-5P**  
610785-45-0P **610785-47-2P** 685877-48-9P 685877-49-0P  
685877-50-3P 685877-51-4P 690958-24-8P 690958-30-6P 690958-38-4P  
**690958-45-3P 690958-80-6P 690958-83-9P**  
**690958-89-5P** 690958-95-3P **690958-98-6P**  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 107724-20-9P, Eplerenone  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 192704-58-8P  
RL: BPN (Biosynthetic preparation); IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 2487-48-1P  
RL: BPN (Biosynthetic preparation); RCT (Reactant); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 54761-04-5, Ytterbium triflate 144026-79-9, Scandium triflate  
RL: CAT (Catalyst use); USES (Uses)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 690959-06-9P 690959-08-1P 690959-11-6P 690959-13-8P  
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)  
(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)
- IT 53-43-0 74-86-2, Acetylene, reactions 534-22-5, 2-Methylfuran  
762-72-1, Allyltrimethylsilane 763-13-3 7677-24-9,



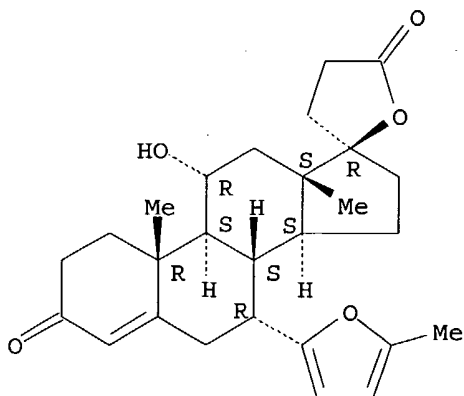
Trimethylsilylcyanide 40474-65-5 67576-47-0 86298-22-8 537718-13-1  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)  
 IT 144-62-7, Oxalic acid, reactions  
 RL: RGT (Reagent); RACT (Reactant or reagent)  
 (preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)  
 IT 75-05-8, Acetonitrile, uses 75-09-2, Dichloromethane, uses  
 RL: NUU (Other use, unclassified); USES (Uses)  
 (solvent; preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)  
 IT 610785-40-5P 610785-47-2P 690958-45-3P  
 690958-80-6P 690958-83-9P 690958-89-5P  
 690958-98-6P  
 RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); RCT  
 (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP  
 (Preparation); RACT (Reactant or reagent)  
 (preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)  
 RN 610785-40-5 HCAPLUS  
 CN Pregna-4,9(11)-diene-21-carboxylic acid, 17-hydroxy-7-(5-methyl-2-furanyl)-  
 3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 610785-47-2 HCAPLUS  
 CN Pregn-4-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

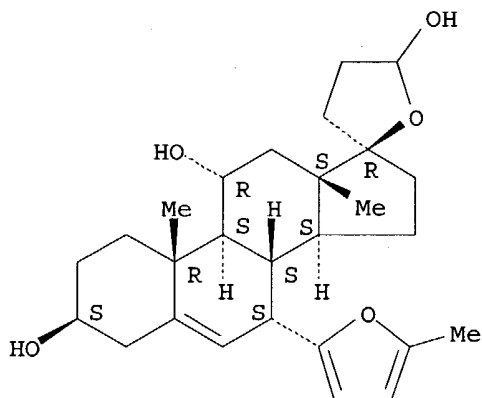
Absolute stereochemistry.



RN 690958-45-3 HCAPLUS

CN Spiro[androst-5-ene-17,2'(3'H)-furan]-3,5',11-triol, 4',5'-dihydro-7-(5-methyl-2-furanyl)-, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )-(9CI) (CA INDEX NAME)

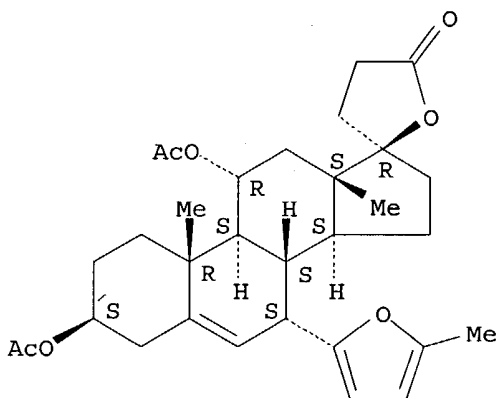
Absolute stereochemistry.



RN 690958-80-6 HCAPLUS

CN Spiro[androst-5-ene-17,2'(5'H)-furan]-5'-one, 3,11-bis(acetyloxy)-3',4'-dihydro-7-(5-methyl-2-furanyl)-, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )-(9CI) (CA INDEX NAME)

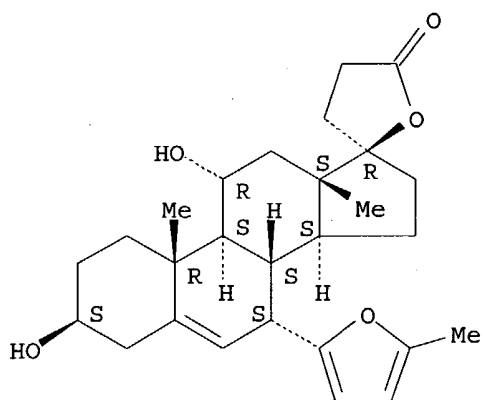
Absolute stereochemistry.



RN 690958-83-9 HCAPLUS

CN Pregn-5-ene-21-carboxylic acid, 3,11,17-trihydroxy-7-(5-methyl-2-furanyl)-,  $\gamma$ -lactone, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )-(9CI) (CA INDEX NAME)

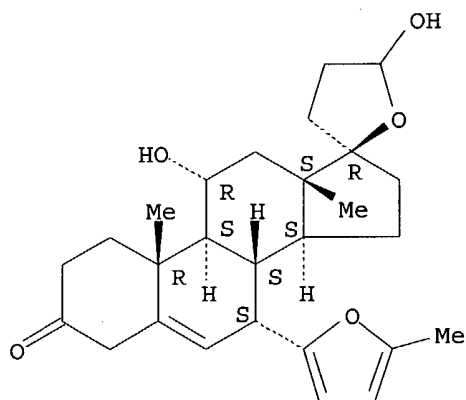
Absolute stereochemistry.



RN 690958-89-5 HCAPLUS

CN Spiro[androst-5-ene-17,2'(3'H)-furan]-3-one, 4',5'-dihydro-5',11-dihydroxy-7-(5-methyl-2-furanyl)-, (7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )-(9CI) (CA INDEX NAME)

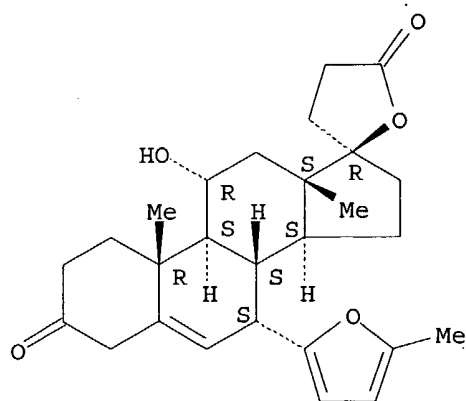
Absolute stereochemistry.



RN 690958-98-6 HCAPLUS

CN Pregn-5-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )-(9CI) (CA INDEX NAME)

Absolute stereochemistry.



L32 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN  
 AN 2003:796726 HCAPLUS  
 DN 139:307925  
 ED Entered STN: 10 Oct 2003  
 TI Process to prepare eplerenone and its intermediates from  
 Δ9-canrenone and other pregnanes  
 IN **Pearlman, Bruce Allen**; Padilla, Amphlett Greg; Havens, Jeffrey  
 L.; Mackey, Sonja S.; Wu, Haifeng  
 PA **Pharmacia & Upjohn Company, USA**  
 SO PCT Int. Appl., 429 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 IC ICM C07J001-00  
 CC 32-5 (Steroids)

Section cross-reference(s): 2, 63

FAN.CNT 2

|      | PATENT NO.      | KIND   | DATE     | APPLICATION NO. | DATE         |
|------|-----------------|--|----------|-----------------|--------------|
| PI   | WO 2003082895   | A2   | 20031009 | WO 2003-US7793  | 20030321 <-- |
|      | WO 2003082895   | A3   | 20040422 |                 |              |
|      | W:              | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |          |                 |              |
|      | RW:             | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |          |                 |              |
|      | US 2003232981   | A1   | 20031218 | US 2003-392833  | 20030321 <-- |
|      | US 2004024202   | A1   | 20040205 | US 2003-392857  | 20030321 <-- |
|      | WO 2004037844   | A1   | 20040506 | WO 2003-US29923 | 20030919 <-- |
|      | WO 2004037844   | C1   | 20040610 |                 |              |
|      | W:              | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD |          |                 |              |
|      | RW:             | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |          |                 |              |
|      | US 2004127702   | A1   | 20040701 | US 2003-666175  | 20030919 <-- |
| PRAI | US 2002-366784P | P  | 20020322 |                 |              |
|      | US 2002-411874P | P  | 20020919 | <--             |              |
|      | US 2002-425596P | P  | 20021112 | <--             |              |
|      | US 2003-392833  | A  | 20030321 |                 |              |

CLASS

PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES

WO 2003082895 ICM C07J001-00  
 OS CASREACT 139:307925; MARPAT 139:307925  
 GI

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB The present invention involves novel intermediates I [R9 = H, OH, O-PG, F; PG = SiMe<sub>3</sub>, SiEt<sub>3</sub>, Ac, CHO; R11 = :O, H<sub>2</sub>, αR11-1βR11-2, R11-5R11-6; R11-1 = H, OR11-3; R11-2 = H, OR11-4; R11-3 = H, PG; R11-4 = H, PG; R11-5R9 = bond, R11-6 = H or R11-6R9 = bond, R11-5 = H; R11-7R9 = O; R11-8 = H; R17 = :O, αR17-1βR17-2, αR17-3βR17-4, αR17-5βR17-6, αR17-7βR17-8, OCH(OR17-9)CH<sub>2</sub>CH<sub>2</sub>, αR17-11βR17-12; R17-1 = H, C.tplbond.CH, CN, C.tplbond.CCH<sub>2</sub>αR17-1-1, C.tplbond.CCH<sub>2</sub>O-PG, CH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>-; R17-2 = OH; R17-3 = OH; R17-4 = COMe, COCH<sub>2</sub>OH, COCH<sub>2</sub>OC(:O)(CH<sub>2</sub>)<sub>0-3</sub>Me; R17-5R17-6 = α-CH<sub>2</sub>O-β; R17-7R17-8 = α-OC(:O)CH<sub>2</sub>CH<sub>2</sub>-β; R17-9 = H, C1-3-alkyl; R17-11 = (CH<sub>2</sub>)<sub>1-2</sub>CH:CH<sub>2</sub>; R17-12 = OH; R17-1-1 = H, Si(R17-1-2)<sub>3</sub>; R17-1-2 = C1-4-alkyl, CH(OEt)Me, THP], including an 7α-substituted steroid, and various novel processes which are used to prepare known intermediates useful in the production of eplerenone, a pharmaceutical agent. Thus, pregnadienone spirolactone II was prepared from Δ<sup>9</sup>-canrenone (III) via conjugate addition of 2-methylfuran in MeNO<sub>2</sub> containing BF<sub>3</sub>·OEt<sub>2</sub>, ring cleavage with dibromantin in aqueous THF containing KOAc, ozonolysis (O<sub>3</sub>/O<sub>2</sub>) in CH<sub>2</sub>Cl<sub>2</sub>/O<sub>2</sub>(CHMe<sub>2</sub>)<sub>2</sub> with Me<sub>2</sub>S quenching in CHCl<sub>3</sub> and oxidation in CHCl<sub>3</sub> with H<sub>2</sub>O<sub>2</sub> in H<sub>2</sub>O containing KHCO<sub>3</sub>.

ST eplerenone steroid prepn; hexahydronaphthalene ketal enol ether prepn

IT Isomerization  
(cis-trans, of pregnenone maleates to fumarates; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Addition reaction  
(conjugate, of furans to pregnenones; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Ethers, reactions  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(enol, alkyl, reaction of, with a hydride abstractor; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Oxidation, electrochemical  
(from alkyl enol ethers; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Abstraction reaction  
(hydrogen, from alkyl enol ethers; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Alcohols, reactions  
Glycols, reactions  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(ketalization by, of Δ<sup>3,5</sup>-dienol ether; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Ozonization  
(of pregnenone maleates and fumarates; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Steroids, preparation  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(pregnanes; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT Ketals  
RL: SPN (Synthetic preparation); PREP (Preparation)  
(Δ<sup>4,6</sup>-; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT 534-22-5, 2-Methylfuran 930-27-8, 3-Methylfuran 7040-43-9, 2-(tert-Butyl)furan  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(conjugate addition of, to pregnadienone derivs.; preparation of eplerenone and its intermediates from Δ<sup>9</sup>-canrenone and other pregnanes)

IT 22037-28-1P, 3-Bromofuran

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
 (conjugate addition of, to pregnadienone derivs.; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 95716-71-5,  $\Delta^9$ -Canrenone 192569-17-8,  $11\alpha$ -Hydroxycanrenone  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (conjugate addition to, by methylfurans; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 192704-56-6,  $11\alpha,17\beta$ -Dihydroxypregn-4-en-3-one  
 $7\alpha,21$ -dicarboxylic acid  $\gamma$ -lactone methyl ester  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (dehydration of; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 67-68-5D, DMSO, o-iodoxybenzoic acid complex 75-91-2D, ruthenium trichloride complex 84-58-2, DDQ 118-75-2, p-Chloranil, reactions 341-02-6, Trityl fluoroborate 536-80-1, Iodosobenzene 546-67-8, Lead tetraacetate 993-02-2, Manganic acetate 1122-96-9D, 4-Methoxypyridine N-oxide, o-iodoxybenzoic acid complex 1184-78-7D, Trimethylamine N-oxide, o-iodoxybenzoic acid complex 1313-13-9, Manganese dioxide, reactions 1333-82-0D, Chromium trioxide, 2,5-dimethylpyrazole complex 2435-53-2, o-Chloranil 2712-78-9, Iodobenzene bistrifluoroacetate 3240-34-4, Iodobenzene diacetate 3375-31-3 7529-22-8D, N-Methylmorpholine N-oxide, o-iodoxybenzoic acid complex 7782-68-5, Iodic acid (HIO<sub>3</sub>) 10049-08-8D, Ruthenium trichloride, tert-Bu hydroperoxide complex 10139-51-2, Ceric ammonium nitrate 12029-98-0, Iodine pentoxide 14546-48-6, Manganese(III), reactions 15158-12-0, Lead+4, reactions 16065-88-6, reactions 18540-29-9, reactions 20492-50-6 22537-44-6, Ruthenium+8, reactions 23317-90-0, Manganese+7, reactions 64297-64-9, o-Iodoxybenzoic acid  
 RL: RGT (Reagent); RACT (Reactant or reagent)  
 (hydrogen abstraction by, in  $\delta^4,6$ -steroid ketals; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 107-21-1, Ethylene glycol, reactions 126-30-7, Neopentyl glycol  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (ketalization by, of  $\Delta^3,5$ -dienol ether; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 192704-66-8  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (ketalization of; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-46-1P  
 RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (preparation and Baeyer-Villiger oxidation of, with peracetic acid; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-42-7P  
 RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (preparation and cis-trans isomerization of; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-55-2P  
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (preparation and crystal structure of; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-47-2P

RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);  
USES (Uses)  
(preparation and dehydration of; preparation of eplerenone and its  
intermediates  
from  $\Delta^9$ -canrenone and other pregnanes)

IT 95716-74-8P  
RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);  
USES (Uses)  
(preparation and esterification or lactone hydrolysis of; preparation of  
eplerenone and its intermediates from  $\Delta^9$ -canrenone and other  
pregnanes)

IT 610785-44-9P  
RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);  
USES (Uses)  
(preparation and intramol. conjugate addition of; preparation of eplerenone  
and its  
intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-45-0P  
RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);  
USES (Uses)  
(preparation and ketalization of; preparation of eplerenone and its  
intermediates  
from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-57-4P  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
(Reactant or reagent)  
(preparation and ozonolysis of; preparation of eplerenone and its  
intermediates  
from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-41-6P 610785-43-8P  
RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);  
USES (Uses)  
(preparation and ozonolysis of; preparation of eplerenone and its  
intermediates  
from  $\Delta^9$ -canrenone and other pregnanes)

IT 610785-50-7P  
RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological  
study); PREP (Preparation); USES (Uses)  
(preparation and ozonolysis of; preparation of eplerenone and its  
intermediates  
from  $\Delta^9$ -canrenone and other pregnanes)

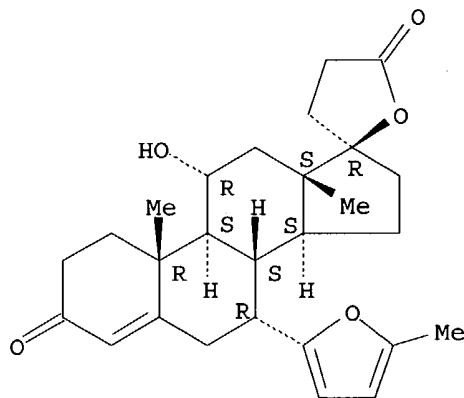
IT 610785-56-3P  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
(Reactant or reagent)  
(preparation and ozonolysis or reaction of, with hydrogen peroxide;  
preparation  
of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other  
pregnanes)

IT 610785-40-5P  
RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);  
USES (Uses)  
(preparation and ring cleavage of, with dibromantin; preparation of  
eplerenone  
and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

IT 107724-20-9P, Eplerenone  
RL: PNU (Preparation, unclassified); PREP (Preparation)  
(preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and

- other pregnanes)
- IT 95717-00-3, 9 $\alpha$ ,11 $\alpha$ -Epoxy-canrenone  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)
- IT 611199-42-9, N-(1,1,2,2,3,3,3-Hexafluoropropyl)diethylamine  
 RL: RGT (Reagent); RACT (Reactant or reagent)  
 (preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)
- IT 95716-70-4P 209253-82-7P 610785-38-1P 610785-39-2P  
 610785-48-3P 610785-49-4P 610785-51-8P  
 610785-52-9P 610785-53-0P 610785-54-1P  
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)
- IT 610785-47-2P  
 RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (preparation and dehydration of; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)
- RN 610785-47-2 HCAPLUS
- CN Pregn-4-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

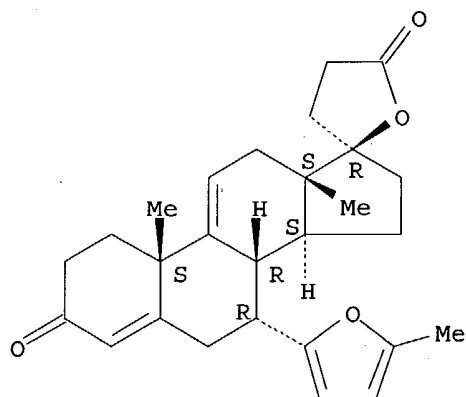
Absolute stereochemistry.



- IT 610785-40-5P  
 RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (preparation and ring cleavage of, with dibromantin; preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)
- RN 610785-40-5 HCAPLUS
- CN Pregn-4,9(11)-diene-21-carboxylic acid, 17-hydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

Absolute stereochemistry.





IT 610785-48-3P 610785-51-8P 610785-52-9P  
610785-53-0P 610785-54-1P

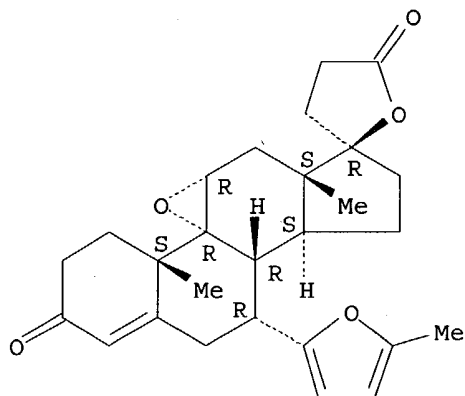
RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of eplerenone and its intermediates from  $\Delta^9$ -canrenone and other pregnanes)

RN 610785-48-3 HCAPLUS

CN Pregn-4-ene-21-carboxylic acid, 9,11-epoxy-17-hydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ ) - (9CI)  
(CA INDEX NAME)

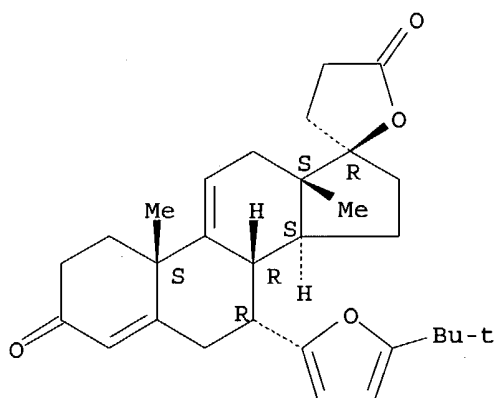
Absolute stereochemistry.



RN 610785-51-8 HCAPLUS

CN Pregna-4,9(11)-diene-21-carboxylic acid, 7-[5-(1,1-dimethylethyl)-2-furanyl]-17-hydroxy-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ ) - (9CI)  
(CA INDEX NAME)

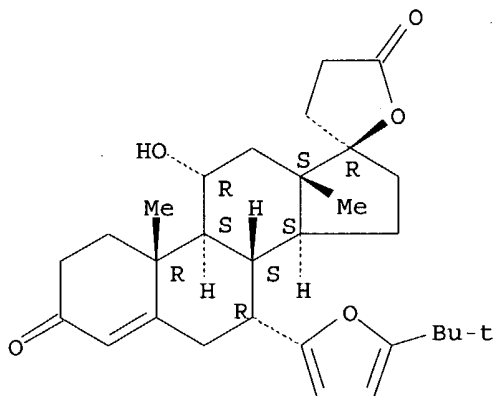
Absolute stereochemistry.



RN 610785-52-9 HCAPLUS

CN Pregn-4-ene-21-carboxylic acid, 7-[5-(1,1-dimethylethyl)-2-furanyl]-11,17-dihydroxy-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI)  
(CA INDEX NAME)

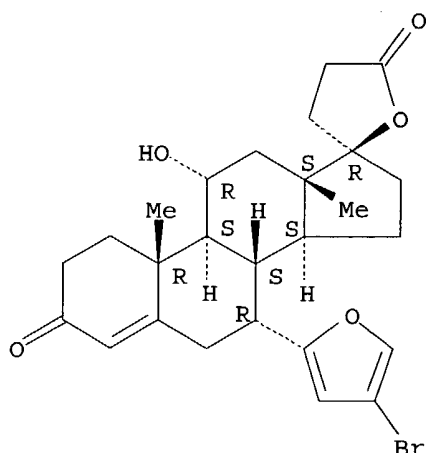
Absolute stereochemistry.



RN 610785-53-0 HCAPLUS

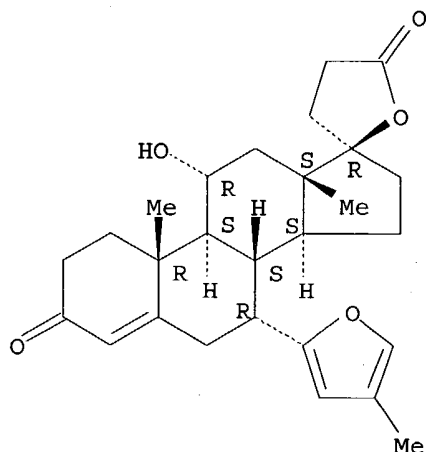
CN Pregn-4-ene-21-carboxylic acid, 7-(4-bromo-2-furanyl)-11,17-dihydroxy-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 610785-54-1 HCAPLUS  
 CN Pregn-4-ene-21-carboxylic acid, 11,17-dihydroxy-7-(4-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )-(9CI) (CA INDEX NAME)

Absolute stereochemistry.



=> fil uspatall

FILE 'USPATFULL' ENTERED AT 06:42:07 ON 28 OCT 2004  
 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 06:42:07 ON 28 OCT 2004  
 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

=> => d l34 bib abs hitstr

L34 ANSWER 1 OF 1 USPATFULL on STN  
 AN 2004:127491 USPATFULL  
 TI Processes for preparing C-7 substituted steroids  
 IN Wuts, Peter Guillaume Marie, Mattawan, MI, UNITED STATES  
 PI US 2004097475 A1 20040520  
 AI US 2003-392945 A1 20030321 (10)  
 PRAI US 2002-424488P 20021107 (60)  
 DT Utility

FS APPLICATION

LREP PHARMACIA & UPJOHN, 301 HENRIETTA ST, 0228-32-LAW, KALAMAZOO, MI, 49007

CLMN Number of Claims: 8

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 975

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention relates to processes for the preparation of novel  
7-carboxy substituted steroid compounds of Formula I, ##STR1##

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

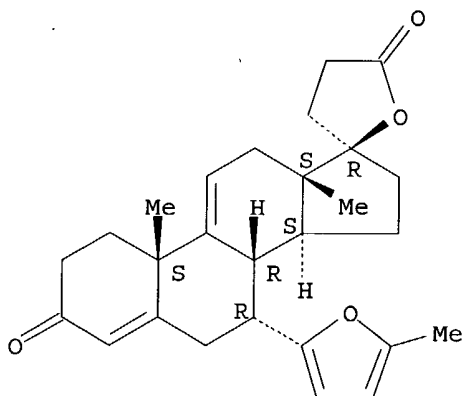
IT 610785-40-5P 610785-47-2P 690958-45-3P  
690958-80-6P 690958-83-9P 690958-89-5P  
690958-98-6P

(preparation of C-7 substituted steroids from 5-androsten-3 $\beta$ -ol-17-one)

RN 610785-40-5 USPATFULL

CN Pregna-4,9(11)-diene-21-carboxylic acid, 17-hydroxy-7-(5-methyl-2-furanyl)-  
3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX NAME)

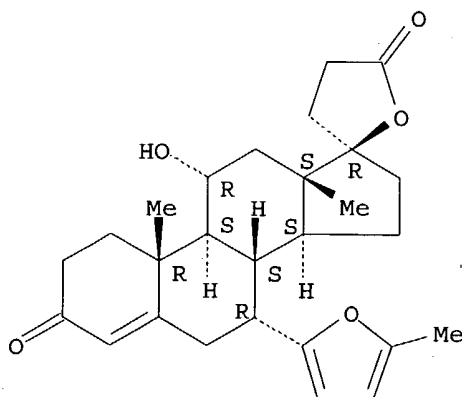
Absolute stereochemistry.



RN 610785-47-2 USPATFULL

CN Pregn-4-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-  
oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )- (9CI) (CA INDEX  
NAME)

Absolute stereochemistry.

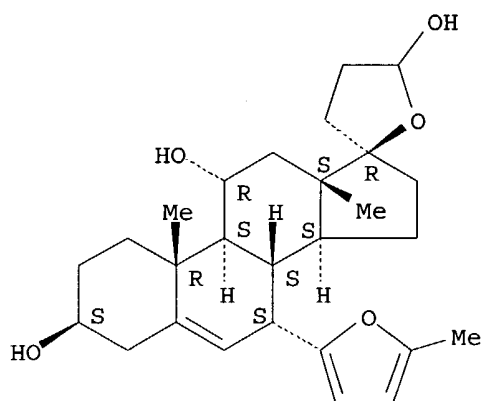


RN 690958-45-3 USPATFULL

CN Spiro[androst-5-ene-17,2'(3'H)-furan]-3,5',11-triol, 4',5'-dihydro-7-(5-  
methyl-2-furanyl)-, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )- (9CI) (CA

INDEX NAME)

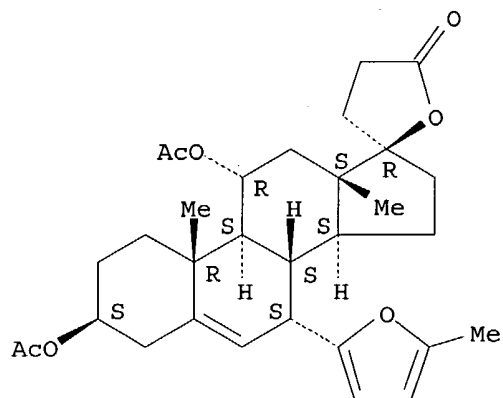
Absolute stereochemistry.



RN 690958-80-6 USPATFULL

CN Spiro[androst-5-ene-17,2' (5'H) -furan] -5'-one, 3,11-bis(acetyloxy) -3',4' -  
dihydro-7-(5-methyl-2-furanyl) -, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ ) -  
(9CI) (CA INDEX NAME)

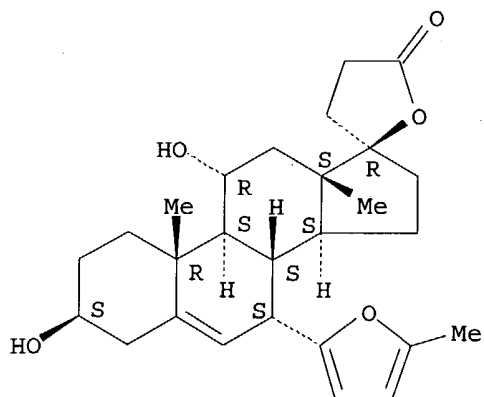
Absolute stereochemistry.



RN 690958-83-9 USPATFULL

CN Pregn-5-ene-21-carboxylic acid, 3,11,17-trihydroxy-7-(5-methyl-2-furanyl) -  
,  $\gamma$ -lactone, (3 $\beta$ ,7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ ) - (9CI) (CA  
INDEX NAME)

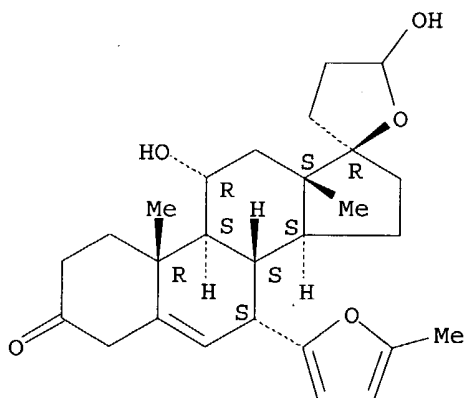
Absolute stereochemistry.



RN 690958-89-5 USPATFULL

CN Spiro[androst-5-ene-17,2' (3'H)-furan]-3-one, 4',5'-dihydro-5',11-dihydroxy-7-(5-methyl-2-furanyl)-, (7 $\alpha$ ,11 $\alpha$ ,17 $\beta$ )-(9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 690958-98-6 USPATFULL

CN Pregn-5-ene-21-carboxylic acid, 11,17-dihydroxy-7-(5-methyl-2-furanyl)-3-oxo-,  $\gamma$ -lactone, (7 $\alpha$ ,11 $\alpha$ ,17 $\alpha$ )-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

